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NEWS		FEB		COMPENDEX reloaded and enhanced
NEWS		FEB		WTEXTILES reloaded and enhanced
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NEWS	20	FEB	23	TOXCENTER updates mirror those of MEDLINE - more
NEWS	21	FEB	23	precise author group fields and 2009 MeSH terms Three million new patent records blast AEROSPACE into
			_	STN patent clusters
NEWS	22	FEB	25	USGENE enhanced with patent family and legal status display data from INPADOCDB
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				equivalents from China

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NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3, AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

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=> s (estradiol or estrogen) and poly(w)D(w)L(w)lactide(w)co(w)glycolide) UNMATCHED RIGHT PARENTHESIS 'GLYCOLIDE)' The number of right parentheses in a query must be equal to the number of left parentheses.

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L2 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2009:109387 CAPLUS

DOCUMENT NUMBER: 150:176325

TITLE: Degradable metal stent with agent-containing coating INVENTOR(S): Klocke, Bjoern; Diener, Tobias; Fringes, Matthias;

Harder, Claus

PATENT ASSIGNEE(S): Biotronik Vi Patent AG, Switz.

SOURCE: Eur. Pat. Appl., 19pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

EP 2018834 A1 20090128 EP 2008-158778 20080623 R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LI, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI,						
SK, TR, AL, BA, MK, RS DE 102007034364 A1 20090129 DE 2007-102007034364 20070724 US 20090030507 A1 20090129 US 2008-179183 20080724 PRIORITY APPLN. INFO:: DE 2007-102007034364A 20070724						
AB The invention concerns a stent comprising: (a) a degradable metal stent main body; (b) a partition layer which is applied to the surface of the stent main body so that at least parts of the surface of the luminal side						
are not covered; (c) an agent-containing layer which is applied to the surface of the partition layer at least partially on the abluminal side of the stent main body, the agent containing a layer comprising one or more agents. The agent comprises at least one polymer selected from the group						
consisting of nondegradable polymers comprising polyethylene; polyvinylchloride; polyacrylates; polyethyl- and polymethylacrylates, polymethylmethacrylate, polymethyl-co-ethyl-acrylate, and						
ethylene/ethylacrylate; polytetrafluoroethylene, ethylene/chlorotrifluoroethylene copolymers, ethylene/tetrafluoroethylene copolymers; polyamides, polyamide imide, PA-11, PA-12, PA-46, PA-66;						
polyetherimide; polyethersulfone; poly(iso)butylene; polyvinylchloride; polyvinylfluoride; polyvinylalc.; polyurethane; polybutylene terephthalate; silicones; polyphosphazene; polymer foams, polymer foams made of carbonates, styrenes; copolymers and blends of the listed polymer						
classes, polymers of the class of thermoplastics, degradable polymers comprising polydioxanone; polyglycolide; polycaprolactone; polylactides, poly-L-lactide, poly-D,L-lactide, and copolymers and blends thereof, poly(L-lactide-co-glycolide), poly(D,L-						
<pre>lactide-co-glycolide), poly(L-lactide-co-D,L-lactide), poly(L-lactide-co-trimethylene carbonate); triblock copolymers; polysaccharides, chitosan, levan, hyaluronic acid, heparin, dextran, cellulose; polyhydroxyvalerate; ethylvinylacetate;</pre>						
polyethylene oxide; polyphosphorylcholine; fibrin; albumin; polyhydroxy butyric acid, atactic, isotactic, and syndiotactic polyhydroxy butyric acid and blends of the foregoing. REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS						

L2 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2009:115360 CAPLUS

DOCUMENT NUMBER: 150:199445

TITLE: Method for production of a crimped stent, use of a

polymer coating and medical goods
Borck, Alexander: Diener, Tobias

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

INVENTOR(S): Borck, Alexander; Diener, Tobias PATENT ASSIGNEE(S): Biotronik Vi Patent A.-G., Switz.

SOURCE: Ger. Offen., 16pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

PATENT NO. APPLICATION NO. DATE KIND DATE ----_____ _____ _____ DE 102007034991 A1 20090129 DE 2007-102007034991 20070726 PRIORITY APPLN. INFO.: DE 2007-102007034991 20070726 The invention concerns a method for the preparation of a crimped stent on a coated catheter by (a) providing a catheter; (b) providing a stent; (c) coating the catheter on the surface that contacts the luminary surface of the stent with a polymer or polymer mixture and one or more active substances; (d) positioning and crimping the stent on the surface of the catheter that has been at least partially coated in step (c). Polymer selected from the group consisting of nondegradable polymers comprising polypropylene, polyethylene; polyvinylchloride; polyacrylates; polyethyland polymethylacrylates, polymethylmethacrylate, polymethyl-co-ethyl-acrylate, and ethylene/ethylacrylate; polytetrafluoroethylene, ethylene/chlorotrifluoroethylene copolymers, ethylene/tetrafluoroethylene copolymers; polyamides, polyamide imide, PA-11, PA-12, PA-46, PA-66; polyether imide; polyethersulfone; poly(iso)butylene; polyvinylchloride; polyvinylfluoride; polyvinylalc.; polyurethane; polybutylene terephthalate; silicones; polyphosphazene; polymer foams, polymer foams made of carbonates, styrenes; copolymers and blends of the listed polymer classes, polymers of the class of thermoplastics, degradable polymers comprising polydioxanone; polyglycolide; polycaprolactone; polylactides, poly-L-lactide, poly-D, L-lactide, and copolymers and blends thereof, poly(L-lactide-co-glycolide), poly(D,Llactide-co-glycolide), poly(L-lactide-co-D,L-lactide), poly(L-lactide-co-trimethylene carbonate); triblock copolymers; polysaccharides, chitosan, levan, hyaluronic acid, heparin, dextran, cellulose; polyhydroxyvalerate; ethylvinylacetate; polyethylene oxide; polyphosphorylcholine; fibrin; albumin; polyhydroxy butyric acid, atactic, isotactic, and syndiotactic polyhydroxy butyric acid and blends of the foregoing. REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 3 OF 10 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: 2008655347 IN-PROCESS

DOCUMENT NUMBER: PubMed ID: 18788777

TITLE: Physicochemical characteristics and preliminary in vivo

biological evaluation of nanocapsules loaded with siRNA

targeting estrogen receptor alpha.

AUTHOR: Bouclier Celine; Moine Laurence; Hillaireau Herve; Marsaud

Veronique; Connault Elisabeth; Opolon Paule; Couvreur

Patrick; Fattal Elias; Renoir Jack-Michel

CORPORATE SOURCE: Physico-Chimie, Pharmacotechnie, Biopharmacie, Universite

Paris-Sud, CNRS UMR 8612 and IFR 141, 5 rue Jean-Baptiste

Clement, 92296 Chatenay-Malabry, France.

SOURCE: Biomacromolecules, (2008 Oct) Vol. 9, No. 10, pp. 2881-90.

Electronic Publication: 2008-09-13.

Journal code: 100892849. E-ISSN: 1526-4602.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

(RESEARCH SUPPORT, NON-U.S. GOV'T)

LANGUAGE: English

FILE SEGMENT: NONMEDLINE; IN-PROCESS; NONINDEXED; Priority Journals

ENTRY DATE: Entered STN: 15 Oct 2008

Last Updated on STN: 1 Jan 2009

AB Specific siRNAs that target estrogen receptor alpha (ERalpha) were encapsulated in nanocapsules (NCs). We produced small (approximately

100-200 nm) ERalpha-siRNA NCs with a water core by incorporating two mixed duplexes of specific ERalpha-siRNAs (ERalpha-mix-siRNA) into NCs. The encapsulation yield that was obtained with poly(iso-butylcyanoacrylate) (PIBCA) NCs was low, whereas no release of trapped siRNA was observed for poly(ethylene)glycol-poly(D,Llactide-co-glycolide) (PEG-PLGA) NCs. High levels of ERalpha-siRNA incorporation into PEG-epsilon-caprolactone-malic acid (PEG-PCL/MA) NCs (3.3 microM in a polymer solution at 16 mg/mL) were observed (72% yield). No difference in size or zeta potential was observed between siRNA NCs that were based on PEG-PCL/MA and empty NCs. Fluorescence quenching assays confirmed the incorporation of siRNA into the NC core. A persistent loss of ERalpha (90% over 5 days) was observed in MCF-7 human breast cancer cells that were exposed to PEG-PCL/MA NCs that were loaded with ERalpha-siRNA. The intravenous injection of these NCs into estradiol-stimulated MCF-7 cell xenografts led to a significant decrease in tumor growth and a decrease in ERalpha expression in tumor cells. These data indicate that a novel strategy, based on ERalpha-siRNA delivery, could be developed for the treatment of hormone-dependent breast cancers.

ANSWER 4 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

2009:165094 CAPLUS ACCESSION NUMBER:

TITLE: Preparation of PLGA microspheres loaded with both

gestodene and ethinyl estradiol

AUTHOR(S): Sun, Yi; Zhang, Qiang; Chen, Da-wei; Zhang, Zhi-jun;

Zheng, Yan

CORPORATE SOURCE: School of Pharmacy, Shenyang Pharmaceutical

University, Shenyang, 110016, Peop. Rep. China

Shenyang Yaoke Daxue Xuebao (2008), 25(12), 948-953 SOURCE:

CODEN: SYDXFF; ISSN: 1006-2858

PUBLISHER: Shenyang Yaoke Daxue Xuebao Bianjibu

DOCUMENT TYPE: Journal LANGUAGE: Chinese

AΒ Objective To study the practicability for the preparation of s.c. injected

microspheres loaded with gestodene and ethinyl estradiol.

Methods Using poly(d, l-lactide-

co-glycolide) (PLGA) as carrier material, gestodene and ethinyl estradiol as model drugs, injected microspheres were prepared through emulsion-evaporation method. The resulted microspheres were observed, and the stability, organic solvent residue and release behavior were also investigated. Results The loading efficiency of gestodene and ethinyl estradiol were (69.9 ± 6.6) % and (60.5 ± 1.5) %, resp. The microspheres were well sphere-shaped, and the particle size distribution was very narrow, and the mean particle size was (65.62 \pm $4.56)\,\mu m$. As a result of the release testing, the release process of both the drugs could last 30 days, and the release behaviors were qualified for Weibull equation, which should be ln(ln1/(1-F(t))) =

 $0.6258 \ln t - 1.826 (R2 = 0.992 1)$ and $\ln (\ln 1/(1-F(t))) = 0.8552 \ln t - 2.8501 (R2 = 0.992 1)$ 0.991 4), resp. The prepared microspheres were not stable when they were

stored at high temperature and lighting condition for a long period, neither

for

long time storage in room temperature, but stable in cool and dark place. Conclusions The preparation condition is stable; the loading efficiency is relatively high; particle size is uniform; the organic residue is qualified for the national standard; release behavior is stable and the process is quite long. The further study should be carried on.

ANSWER 5 OF 10 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2007:949868 CAPLUS

DOCUMENT NUMBER: 147:474420

TITLE: Design of surface-modified poly(D,

L-lactide-co-

glycolide) nanoparticles for targeted drug

delivery to bone

AUTHOR(S): Choi, Sung-Wook; Kim, Jung-Hyun

CORPORATE SOURCE: Nanosphere Process and Technology Laboratory,

Department of Chemical Engineering, Yonsei University,

Seoul, 120-749, S. Korea

SOURCE: Journal of Controlled Release (2007), 122(1), 24-30

CODEN: JCREEC; ISSN: 0168-3659

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Poly(D,L-lactide-co-

glycolide) (PLGA) nanoparticles, modified with both alendronate and polyethylene glycol (PEG), were prepared by dialysis method without addnl. surfactant to evaluate the potency of the bone-targeted drug delivery. Alendronate, a targeting moiety that has a strong affinity for bone, was conjugated to PLGA polymer via carbodiimide chemical Monomethoxy PEG(mPEG)-PLGA block copolymers with different mol. wts. of mPEG (Mn 550, 750, and 2000) were synthesized and used for a hydrophilic layer on the surface of the nanoparticles to avoid reticuloendothelial system (RES). The surface-modified PLGA nanoparticles with various ratios of alendronate and mPEG densities on their surface were evaluated by adsorption study onto hydroxyapatite (HA). It was confirmed that alendronate-modified nanoparticles had a strong and specific adsorption to HA. The amount of nanoparticles absorbed onto HA tended to be smaller when the content of alendronate was decreased and the large block length of mPEG was found to reduce the potency of alendronate.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 6 OF 10 MEDLINE on STN DUPLICATE 2

ACCESSION NUMBER: 2004056339 MEDLINE DOCUMENT NUMBER: PubMed ID: 14757510

TITLE: Pure antiestrogen RU 58668-loaded nanospheres: morphology,

cell activity and toxicity studies.

AUTHOR: Ameller Thibault; Marsaud Veronique; Legrand Philippe; Gref

Ruxandra; Renoir Jack Michel

CORPORATE SOURCE: UMR CNRS 8612, Pharmacologie Cellulaire et Moleculaire, 5

rue Jean-Baptiste Clement, 92296, Chatenay-Malabry,

France.. michael.renoir@cep.u-psud.fr

SOURCE: European journal of pharmaceutical sciences: official

journal of the European Federation for Pharmaceutical Sciences, (2004 Feb) Vol. 21, No. 2-3, pp. 361-70.

Journal code: 9317982. ISSN: 0928-0987.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: (COMPARATIVE STUDY)

Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, NON-U.S. GOV'T)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200411

ENTRY DATE: Entered STN: 4 Feb 2004

Last Updated on STN: 19 Dec 2004 Entered Medline: 26 Nov 2004

AB Nanospheres (NS) formulated using biodegradable and biocompatible

polymers, poly(D,L-lactide-

co-glycolide) (PLGA), poly(D,L-lactide) (PLA) and

poly(epsilon-caprolactone) (PCL), loaded with the pure anti-

estrogen RU 58668 (RU), a promising estrogen-dependent

anticancer agent, have been prepared. They all possess a small size compatible with an intratumoral extravasation behavior and their

pegylation reduce significantly their zeta potential. Characterization by

freeze fracture electron microscopy have shown that NS are spheric particles with a size ranging between 30 and 50nm and a tendency to agglomerate which is reduced by polyethylene glycol (PEG) grafting. PEG-grafted NS are all non-toxic as revealed by cell viability assay. A specific cellular model has been used to evaluate not only the release extent of the drug but also its biological activity. All formulations tested showed that they release slowly RU as measured by the delayed ability of RU to inhibit estrogen-induced transcription in human breast cancer cells and that they possess only a small amount of surface adsorbed RU.

L2 ANSWER 7 OF 10 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2001104067 EMBASE

TITLE: Preparation of PLGA nanoparticles containing estrogen by emulsification-diffusion method.

AUTHOR: Kwon, Hye-Young; Lee, Jun-Young; Choi, Sung-Wook; Kim,

Jung-Hyun (correspondence)

CORPORATE SOURCE: Nanosphere Process and Technology Laboratory, Department of

Chemical Engineering, Yonsei University, 134 Shinchon-dong, Sudaemoon-ku Seoul 120-749, Korea, Republic of. jayhkim@mai

1.yonsei.ac.kr

AUTHOR: Jang, Yangsoo

CORPORATE SOURCE: Yonsei University, 134 Shinchon-dong, Sudaemoon-ku, Seoul

120749, Korea, Republic of.

AUTHOR: Kim, Jung-Hyun (correspondence)

CORPORATE SOURCE: Nanosphere Process/Technol Lab, Department of Chemical

Eng., Yonsei University, 134 Shinchon-dong, Sudaemoon-ku, Seoul 120-749, Korea, Republic of. jayhkim@mail.yonsei.ac.k

r

SOURCE: Colloids and Surfaces A: Physicochemical and Engineering

Aspects, (30 Jun 2001) Vol. 182, No. 1-3, pp. 123-130.

Refs: 19

ISSN: 0927-7757 CODEN: CPEAEH

PUBLISHER IDENT.: S 0927-7757(00)00825-6

COUNTRY: Netherlands
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 027 Biophysics, Bioengineering and Medical

Instrumentation

037 Drug Literature Index

039 Pharmacy

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 6 Apr 2001

Last Updated on STN: 6 Apr 2001

AB Nano-sized poly (D,L lactide-

co-glycolide) (PLGA) particles, widely used as a biodegradable polymeric carrier, containing estrogen were prepared employing emulsification-diffusion method. Estrogen was chosen as a model drug. The preparation method consists of emulsifying a solution of polymer and drug in the aqueous phase containing stabilizer, previously saturated, followed by adding excess water. Influence of process variables on the mean particle size of nanoparticles has been studied. It was clarified that the type and concentrations of stabilizer, homogenizer speed, polymer concentration determined the size of PLGA nanoparticles. Especially when didodecyl dimethyl ammonium

bromide (DMAB) was used as a stabilizer, estrogen containing nanoparticles of smaller than 100 nm was obtained. Copyright .COPYRGT. 2001 Elsevier Science B.V.

L2 ANSWER 8 OF 10 MEDLINE on STN ACCESSION NUMBER: 1999401151 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10469916

TITLE: Gamma irradiation for terminal sterilization of 17beta-

estradiol loaded poly-(D, L-lactide-co-glycolide

) microparticles.

AUTHOR: Mohr D; Wolff M; Kissel T

CORPORATE SOURCE: Schwarz Pharma AG, D-40789, Monheim, Germany.

SOURCE: Journal of controlled release: official journal of the Controlled Release Society, (1999 Aug 27) Vol. 61, No. 1-2,

pp. 203-17.

Journal code: 8607908. ISSN: 0168-3659.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199910

ENTRY DATE: Entered STN: 26 Oct 1999

Last Updated on STN: 26 Oct 1999 Entered Medline: 13 Oct 1999

AB 17beta-Estradiol-loaded microparticles using poly-(

D, L-lactide-co-glycolide)

polymer (PLG) were prepared by a modified spray-drying method and the effects of gamma-irradiation on drug substance, polymer and microparticles were investigated. Irradiation doses ranging from 5.1 to 26.6 kGy were applied using a 60Co-radiation source. 17beta-Estradiol drug substance showed excellent stability against gamma-irradiation in the investigated dose range, whereas microencapsulated estradiol seems to be converted to conjugation products with PLG, and to a lesser extent to the degradation product 9,11-dehydroestradiol. The weight-average molecular weight of the PLG polymers decreased with increasing irradiation dose while polydispersity indices (M(w)/M(n))remained nearly unchanged, compatible with a random chain scission mechanism in lactide/glycolide-copolymer degradation. In vitro drug release studies showed accelerated kinetics with increasing irradiation doses due to dose dependent polymer degradation. Microbiological process monitoring showed decreasing bioburden with increasing spraying time, which was successfully further reduced by applying irradiation sterilization. Microencapsulated test spore suspensions of Bacillus pumilus ATCC 27142, the official test specimen for the gamma-sterilization process, revealed effective reduction of bioburden, confirming its published D(10) value. In conclusion, our studies demonstrated efficacy of gamma-irradiation as terminal sterilization method for poly-(D, L-lactide-co-glycolide)

polymer-based drug delivery systems. The sterilization conditions need to be carefully adjusted for the final dosage form.

L2 ANSWER 9 OF 10 MEDLINE on STN ACCESSION NUMBER: 1994007813 MEDLINE DOCUMENT NUMBER: PubMed ID: 8403904

TITLE: The effects of a long-acting progestin on the

hypothalamic-pituitary-ovarian axis in women with normal

menstrual cycles.

AUTHOR: Poindexter A N 3rd; Dildy G A; Brody S A; Snabes M C;

Brodyand S A

CORPORATE SOURCE: Department of Obstetrics and Gynecology, Baylor College of

Medicine, Houston, TX.

CONTRACT NUMBER: MOIRR000350 (United States NCRR NIH HHS)

SOURCE: Contraception, (1993 Jul) Vol. 48, No. 1, pp. 37-45.

Journal code: 0234361. ISSN: 0010-7824.

PUB. COUNTRY: United States
DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RESEARCH SUPPORT, NON-U.S. GOV'T) (RESEARCH SUPPORT, U.S. GOV'T, P.H.S.)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199310

ENTRY DATE: Entered STN: 17 Jan 1994

Last Updated on STN: 25 Feb 2003 Entered Medline: 28 Oct 1993

AB This study was performed to determine how a long-acting, slow-release preparation of norethindrone (NET) affects the hypothalamic-pituitary-ovarian axis of normal ovulatory women. Ten women were studied during the luteal phase of their menstrual cycle, and again at six and twelve weeks following intramuscular administration of 100 mg NET microencapsulated in poly-D,L-lactide-co-glycolide. Serial LH samples,

serum E, P, and NET were followed by a GnRH stimulation test. Compared to luteal phase values, six and twelve weeks of treatment with NET inhibited serum E2 and P while mean serum LH remained unchanged and mean serum FSH increased significantly (p < 0.05). LH pulse frequency after NET treatment was twice the rate (p < 0.01) as that of the luteal phase, whereas LH pulse amplitude was decreased significantly (p < 0.05). Finally, although there was no significant change in pituitary LH secretion in response to GnRH, NET treatment augmented FSH responsiveness to GnRH at the times studied. Preserved pituitary responsiveness to GnRH in NET-treated patients suggests that inhibited ovarian function results in an increase in GnRH pulse frequency but not GnRH pulse amplitude. Since the progestational milieu is maintained in these patients by NET treatment, the decrease in serum E2 may be responsible for the increase in GnRH pulse frequency. The presence of a critical level of E2 may be necessary for progestins to affect the hypothalamic GnRH pulse generator.

L2 ANSWER 10 OF 10 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 4

ACCESSION NUMBER: 1983:332927 BIOSIS

DOCUMENT NUMBER: PREV198376090419; BA76:90419

TITLE: POLY-D L LACTIDE

CO GLYCOLIDE NORETHISTERONE MICRO

CAPSULES AN INJECTABLE BIO DEGRADABLE CONTRACEPTIVE.

AUTHOR(S): BECK L R [Reprint author]; POPE V Z; FLOWERS C E JR; COWSAR

D R; TICE T R; LEWIS D H; DUNN R L; MOORE A B; GILLEY R M

CORPORATE SOURCE: DEPARTMENT OF OBSTETRICS AND GYNECOLOGY, DIVISION FOR

REPRODUCTION AND IMMUNOLOGY RESEARCH, UNIVERSITY OF ALABAMA

IN BIRMINGHAM, SCHOOL OF MEDICINE, BIRMINGHAM, ALABAMA

35294, USA

SOURCE: Biology of Reproduction, (1983) Vol. 28, No. 1, pp.

186-195.

CODEN: BIREBV. ISSN: 0006-3363.

DOCUMENT TYPE: Article FILE SEGMENT: BA LANGUAGE: ENGLISH

AB Microcapsules made from a biocompatible, biodegradable polymeric excipient, poly(DL-lactide-co-glycolide) (DL-PLGA) that contained 22 wt % norethisterone (NET), were prepared by a solvent-evaporation microencapsulation process. The effects of changing both the lactide-to-glycolide ratio of the DL-PLGA and the size of the microcapsules on the rate of NET release and the rate of excipient biodegradation were determined in vivo. NET release rates were determined in baboons after injecting the microcapsule formulations i.m. Serum samples obtained at various times following treatment were analyzed for NET, progesterone and estrogen by radioimmunoassay (RIA). Biodegradation kinetics were determined by injecting NET microcapsules made from radiolabeled DL-PLGA i.m. into the hind legs of rats. Residual

radioactivity at the injection site was determined at various times after treatment by combustion analysis of the muscle tissue. Changing the ratio of the comonomers to include more glycolide (DL-lactide:glycolide 96:4, 92:8, 87:13, 74:26) increased the rate of NET release and accelerated the biodegradation of the copolymer excipient. Decreasing the size of the microcapsules increased the rate of NET release. On the basis of these studies a NET microcapsule formulation was identified for clinical testing which releases NET for 3 mo. and biodegrades completely within 6 mo.

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